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The following is a detailed listing of all claims that are, or were, pending in the present application. Please amend claims 32 and 66, as set forth in this detailed listing.

**Detailed Claim Listing** 

Claims 1-23 (canceled)

- 24. (previously presented): A composition comprising:
- a) a substrate with a surface comprising discrete sites at a density of at least 100 discrete sites per 1 mm<sup>2</sup>, said discrete sites comprising wells; and
- b) a population of microspheres randomly distributed in said wells, said population comprising at least a first and a second subpopulation, said microspheres comprising a bioactive agent, and wherein said sites can have only a single microsphere.
  - 25. (previously presented): A composition comprising:
- a) a substrate with a patterned surface comprising discrete sites, said substrate comprising discrete sites at a density of at least 100 discrete sites per 1 mm<sup>2</sup>; and
- b) a population of microspheres, randomly distributed on said sites, wherein each microsphere comprises a bioactive agent; and

wherein said sites can have only a single microsphere.

- 26. (previously presented): A composition according to claim 24 or 25 wherein said substrate is a fiber optic bundle.
- 27. (previously presented): A composition according to claim 24 or 25 wherein said substrate is selected from the group consisting of glass and plastic.
- 28. (previously presented): A composition according to claim 24 wherein said population of microspheres comprises at least a first and a second subpopulation, wherein the microspheres of said first subpopulation of microspheres are a different size than the microspheres of said second subpopulation.

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29. A composition according to claim 24 or 25 wherein (previously presented): said bioactive agent comprises a protein.

- 30. (previously presented): A composition according to claim 29 wherein said protein is selected from the group consisting of enzymes and antibodies.
- 31. (previously presented): A composition according to claim 24 or 25 wherein said bioactive agent is a nucleic acid.
- (currently amended): A composition according to claim 66-wherein said 32. population of microspheres comprises at least a first and a second subpopulation, wherein the microspheres of said first subpopulation of microspheres are a different size than the microspheres of said second subpopulation.
- A composition according to claim 24, 66, 28, or 32 33. (previously presented): wherein said first and said second subpopulations comprise a first and a second bioactive agent, respectively.
- The composition according to claim 33, wherein 34. (previously presented): said first and second subpopulations further comprise a first and a second optical signature, respectively.
- A composition according to claim 34 wherein said 35. (previously presented): at least one of said optical signatures comprises at least one chromophore.
- (previously presented): 36. A composition according to claim 34 wherein said at least one of said optical signatures comprises at least one fluorescent dye.

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37. (previously presented): A composition according to claim 36 wherein said fluorescent dye is entrapped within said microspheres.

- 38. (previously presented): A composition according to claim 36 wherein said fluorescent dye is attached to said microspheres.
- 39. (previously presented): A composition according to claim 34 wherein said optical signature comprises at least two fluorescent dyes.
- 40. (previously presented): A composition according to claim 66 wherein said bioactive agent comprises a protein.
- 41. (previously presented): A composition according to claim 40 wherein said protein is selected from the group consisting of enzymes and antibodies.
- 42. (previously presented): A composition according to claim 66 wherein said bioactive agent is a nucleic acid.
- 43. (previously presented): A composition according to claim 24 or 25 wherein said bead is covalently associated with the well.
- 44. (previously presented): A composition according to claim 24 or 25 wherein said bead is non-covalently associated with the well.
- 45. (previously presented): A method of determining the presence of at least a first and second target analyte in a sample comprising:
  - a) contacting said sample with a composition comprising:
    - i) a substrate with a patterned surface comprising discrete sites; and
    - ii) a population of microspheres comprising at least a first and a second subpopulation, wherein said first subpopulation comprises a first bioactive

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agent and said second subpopulation comprises a second bioactive agent, wherein said microspheres are randomly distributed on said surface such that said discrete sites contain only one microsphere; and

- b) determining the presence of said first and second target analyte.
- 46. (previously presented): A method according to claim 45 wherein said substrate is a optical fiber bundle and said microspheres are located within wells at a first terminal end of said bundle.
- 47. (previously presented): A method according to claim 45 further comprising identifying the location of said first and second bioactive agent on said substrate.
- 48. (previously presented): The method according to claim 45, wherein said discrete sites are wells.
- 49. (previously presented): The method according to claim 45, wherein said substrate is selected from the group consisting of glass and plastic.
  - 50. (previously presented): A method of making a composition comprising:
  - a) providing a patterned surface comprising individual sites on a substrate;
- b) randomly distributing microspheres on said surface such that said individual sites contain microspheres, wherein said sites can have only a single microsphere, and wherein said microspheres comprise at least a first and a second subpopulation comprising:
  - i) a first and second bioactive agent, respectively; and
  - ii) a first and second optical signature, respectively;
- c) detecting said first and second optical signatures while said microspheres are distributed on said surface; and
- d) correlating the location of at least one individual site on the array with the bioactive agent at that particular site.

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51. (previously presented): A method according to claim 50, wherein said distributing comprises serially adding said subpopulations to said sites.

- 52. (previously presented): A method according to claim 50, wherein said substrate is a fiber optic bundle.
- 53. (previously presented): A method according to claim 50, wherein said substrate is selected from the group consisting of glass and plastic.
- 54. (previously presented): A method according to claim 50, wherein said sites are wells.
- 55. (previously presented): A method according to claim 45 or 50, wherein said bead is covalently attached to the well.
- 56. (previously presented): A method according to claim 45 or 50, wherein said bead is non-covalently attached to the well.
- 57. (previously presented): A method according to claim 45 or 50, wherein said bioactive agent is a nucleic acid.
- 58. (previously presented): A composition according to claim 27 wherein said substrate is glass.
- 59. (previously presented): A composition according to claim 27 wherein said substrate is plastic.
- 60. (previously presented): A composition according to claim 30 wherein said protein is an enzyme.

61. (previously presented): A composition according to claim 30 wherein said protein is an antibody.

- 62. (previously presented): A composition according to claim 41 wherein said protein is an enzyme.
- 63. (previously presented): A composition according to claim 41 wherein said protein is an antibody.
- 64. (previously presented): A method according to claim 49 or 53 wherein said substrate is glass.
- 65. (previously presented): A method according to claim 49 or 53 wherein said substrate is plastic.
- 66. (currently amended): A <u>method</u>composition according to claim 25, wherein said population of microspheres comprises at least a first and a second subpopulation.
- 67. (previously presented): A method according to claim 45 or 50 when said bioactive agent is a protein.